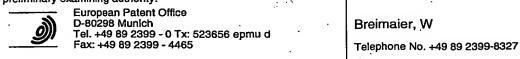
***** PATENT COOPERATION TREATY**

PCT

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

		(PCT Article 36	and	Rule 70)			
						REC'D	1 1 OCT 2	2004
				_		WIPO	P	СТ
Applicant's SCB 762		ent's file reference	FOR FURTHER A	CTION	See Notification Preliminary Exa			
Internationa	• •		International filing date	(day/mon	th/year)	Priority date	e (day/month/)	year)
PCT/EP (15.04.2003					·
C07C49/		nt Classification (IPC) or bot	n national classification :	and IPC				
Applicant INDENA	S D	A ot al				-		
INDENA	5.F. <i>i</i>							
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1. This Auth	interi ority	national preliminary exam and is transmitted to the a	ination report has bee applicant according to	en prepa Article 3	red by this inter 36.	national Pr	əliminary Ex	camining
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2. This	REP	ORT consists of a total of	4 sheets, including t	his cove	r sheet.			
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	bee	report is also accompant n amended and are the ba Rule 70.16 and Section	asis for this report and	d/or shee	ets containing re	ctifications	made befor	e this Authorit
The	•	nexes consist of a total of				,		:
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3. This		rt contains indications rela	ating to the following i	tems;		,		
1	Ø	Basis of the opinion					•	
li III		Priority	minion with record to	aavaltu i	inventive eten e	nd industric	al applicabili	to e
111		Non-establishment of o	_	loveity, i	inventive step at	iid iiidusiiid	ıı applicabili	ty
IV V		Lack of unity of invention Reasoned statement un		iith rogai	rd to povelty in	ontiva ster	or industris	al applicability:
V		citations and explanation	ons supporting such st	tatement	t	entive step	, or madsum	ar applicability,
VI		Certain documents cite	d					
VII		Certain defects in the ir	nternational application	n				
VIII		Certain observations or	n the international app	lication				
Date of sul	omissi	on of the demand		Date o	of completion of the	s report		
12.11.20	03			08.10	0.2004			
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INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.

PCT/EP 03/03923

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1. With regard to the **elements** of the international application (Replacement sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to this report since they do not contain amendments (Rules 70.16 and 70.17)):

	Des	cription, Pages					
	1-13	3	as originally filed				
	Olai	ima Numbana	·				
		ims, Numbers					
	1-7		as originally filed				
2.	With lang	Vith regard to the language , all the elements marked above were available or furnished to this Authority in the anguage in which the international application was filed, unless otherwise indicated under this item.					
	The	se elements were ava	ailable or furnished to this Authority in the following language: , which is:				
		the language of a tra	nslation furnished for the purposes of the international search (under Rule 23.1(b)).				
		the language of publ	ication of the international application (under Rule 48.3(b)).				
		the language of a tra Rule 55.2 and/or 55.3	nslation furnished for the purposes of international preliminary examination (under 3).				
3.	Witl inte	With regard to any nucleotide and/or amino acid sequence disclosed in the international application, the international preliminary examination was carried out on the basis of the sequence listing:					
		contained in the inter	national application in written form.				
		filed together with the	e international application in computer readable form.				
		furnished subsequer	tly to this Authority in written form.				
		furnished subsequen	tly to this Authority in computer readable form.				
		The statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.					
		The statement that the listing has been furnited	ne information recorded in computer readable form is identical to the written sequence shed.				
4.	The	amendments have re	esulted in the cancellation of:				
		the description,	pages:				
		the claims,	Nos.:				
		the drawings,	sheets:				
5.			established as if (some of) the amendments had not been made, since they have go beyond the disclosure as filed (Rule 70.2(c)).				
		(Any replacement sh report.)	eet containing such amendments must be referred to under item 1 and annexed to this				
6.	Add	litional observations, i	f necessary:				

INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.

PCT/EP 03/03923

- V. Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement
- 1. Statement

Novelty (N) Yes: Claims 1-7

No: Claims

Inventive step (IS) Yes: Claims 1-7

No: Claims

Industrial applicability (IA) Yes: Claims 1-7

No: Claims

2. Citations and explanations

see separate sheet

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Re Item V

Reasoned statement with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

D1: WO 99/64388 D2: WO 99/41220

The present invention concerns compounds of general formula (I) which are said to possess anti-depressant, anxiolytic and anti-neurodegenrative activities according to claims 1 to 7.

novelty

The subject-matter according to claims 1 to 7 is novel pursuant to Art. 33(2) PCT.

D1 describes OH-functionalized (ad)hyperforin derivatives of general formula (I) as an anti-depressant (see pages 5 and 6 and the claims).

D2 describes stable (ad)hyperforin salts of general formula (I) which are useful as antidepressants and for treating eg Alzheimer's disease (see page 6, tables II-IV and claims).

None of the documents of the available prior art describes the 8-halogen substituted derivatives of (ad)hyperforin of general formula (I) according to claim 1.

Thus, novelty for the subject-matter according to claims 1 to 7 is given.

inventive step

The subject-matter according to claims 1 to 7 is inventive pursuant to Art. 33(3) PCT.

In view of the closest state of the art D1 and D2, the problem posed is the provision of further stable (ad)hyperforin derivatives which possess anti-depressant and anti-neurodegenerative activities.

The above problem is solved by (ad)hyperforin derivatives which are substituted at the 8-position with halogen according to claim 1. The results reported on page 8 show that the halogenated compounds of the invention possess the desired activities. In particular, the compounds as claimed are more active than the parent hyperforin. In the absence of any indication in the available prior art that substitution of (ad)hyperforin with halogen at the 8-position would furnish the corresponding stable derivatives with the desired activity, an inventive step is acknowledged.